**Supplementary Figure S1.** Mobility of Dronpa-θ-D4 molecules. (A) Diffraction limited image of cholesterol enriched domains integrated over 1000 frames. Scale bar: 5 µm. (B) Distribution of the diffusion coefficients of trajectories described by Dronpa-θ-D4 molecules that remain fluorescent for more than 4 consecutive frames (corresponding to 124 ms). The diffusion coefficient is calculated using the first 3 points of the MSD for each individual trajectory (for a total of 220 trajectories). The mean value of the diffusion coefficient (0.013 µm²/s) corresponds to a mean displacement of 46 nm between consecutive frames. (C) Trajectories described by Dronpa-θ-4 molecules in the area indicated in (A). Each color indicates a different track. Although mobile, the molecules are confined within the cholesterol enriched domains. (D) Ten randomly chosen trajectories present in the domain indicated by the black square in (C). Each Dronpa-θ-4 molecule covers an area smaller than the total area of the domain.